

**Listing of Claims**

Please amend claims 1, 4, 29, 34-36, 38 and 41, and please add new claims 43-48, as shown below. This listing of claims will replace all prior versions and listings of claims in the instant application.

1. (Currently Amended) An antimicrobial sulfonamide derivative, or a salt or a hydrate thereof, comprising:

a core cyclic peptide or core antibiotic of ~~a~~ an acidic lipopeptide antibiotic; and  
a lipophilic moiety,

wherein said lipophilic moiety is covalently attached to the core cyclic peptide or core antibiotic *via* a linking chain which includes a sulfonamide linkage and wherein said core cyclic peptide or core antibiotic is not of laspartomycin ~~or polymyxin~~.

2. (Original) The antimicrobial sulfonamide derivative, salt or hydrate of Claim 1 in which the linking chain is a sulfonamide linkage.

3. (Original) The antimicrobial sulfonamide derivative, salt or hydrate of Claim 1 in which the linking chain is a linker that links the core cyclic peptide or core antibiotic to the lipophilic moiety.

4. (Currently Amended) The antimicrobial sulfonamide derivative, salt or hydrate of Claim 1 which is a compound according to structural Formula (I):

(I)  $Y-X-N(R^4)(-L-X-N(R^1))_m-R$

wherein:

Y is a lipophilic moiety;

each X is independently selected from the group consisting of ~~—~~ $\text{SO}_2^-$ ~~—~~ $\text{CO-SO}_2^-$ , ~~—~~ $\text{CS}$ ~~—~~, ~~—~~ $\text{PO}$ ~~—~~, ~~—~~ $\text{OP(O)}$ ~~—~~, ~~—~~ $\text{OC(O)}$ ~~—~~, ~~—~~ $\text{NHCO}$ ~~—~~ and ~~—~~ $\text{N}(R^1)\text{CO}$ ~~—~~ with the proviso that at least one X is ~~—~~ $\text{SO}_2^-$ ~~—~~ $\text{SO}_2^-$ ;

M is 0 or 1;

L is a linker;

N is nitrogen;

R<sup>1</sup> and R<sup>4</sup> are each independently selected from the group consisting of hydrogen, (C<sub>1</sub>-C<sub>25</sub>) alkyl optionally substituted with one or more of the same or different R<sup>2</sup> groups, (C<sub>1</sub>-C<sub>25</sub>) heteroalkyl optionally substituted with one or more of the same or different R<sup>2</sup> groups, (C<sub>5</sub>-C<sub>30</sub>) aryl optionally substituted with one or more of the same or different R<sup>2</sup> groups, (C<sub>5</sub>-C<sub>30</sub>) arylaryl optionally substituted with one or more of the same or different R<sup>2</sup> groups, (C<sub>5</sub>-C<sub>30</sub>) biaryl optionally substituted with one or more of the same or different R<sup>2</sup> groups, five to thirty membered heteroaryl optionally substituted with one or more of the same or different R<sup>2</sup> groups, (C<sub>6</sub>-C<sub>30</sub>) arylalkyl optionally substituted with one or more of the same or different R<sub>2</sub> groups and six to thirty membered heteroarylalkyl optionally substituted with one or more of the same or different R<sub>2</sub> groups;

each R<sup>2</sup> is independently selected from the group consisting of -OR<sup>3</sup>, -SR<sup>3</sup>, -NR<sup>3</sup>R<sup>3</sup>, -CN, -NO<sub>2</sub>, -N<sup>3</sup>-N<sub>3</sub>, -C(O)OR<sup>3</sup>, -C(O)NR<sup>3</sup>R<sup>3</sup>, -C(S)NR<sup>3</sup>R<sup>3</sup>, -C(NR<sup>3</sup>)NR<sup>3</sup>R<sup>3</sup>, -CHO, -R<sup>3</sup>CO, -SO<sub>2</sub>R<sup>3</sup>, -SOR<sup>3</sup>, -PO(OR<sup>3</sup>)<sub>2</sub>, -PO(OR<sup>3</sup>), -CO<sub>2</sub>H, -SO<sub>3</sub>H, -PO<sub>3</sub>H, halogen and trihalomethyl;

each R<sup>3</sup> is independently selected from the group consisting of hydrogen, (C<sub>1</sub>-C<sub>6</sub>) alkyl, (C<sub>5</sub>-C<sub>10</sub>) aryl, five to sixteen membered heteroaryl, (C<sub>6</sub>-C<sub>16</sub>) arylalkyl and six to sixteen membered heteroarylalkyl; and

R is a core cyclic peptide or core antibiotic of a an acidic lipopeptide antibiotic, wherein said core cyclic peptide or core antibiotic is not of laspartomycin or polymyxin.

5. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core cyclic peptide of zaomycin, crystallomycin, aspartocin, amphotomycin, glumamycin, brevistin, cerexin A, cerexin B, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A-54145, Antibiotic A-21978C or tsushimycin.

6. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core antibiotic of zaomycin, crystallomycin, aspartocin, amphotomycin, glumamycin, brevistin, cerexin A, cerexin B, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A-54145, Antibiotic A-21978C or tsushimycin.

7. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core cyclic peptide of aspartocin, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A-54145 or Antibiotic A-21978C.

8. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core antibiotic of aspartocin, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A54145 or Antibiotic A-21978C.

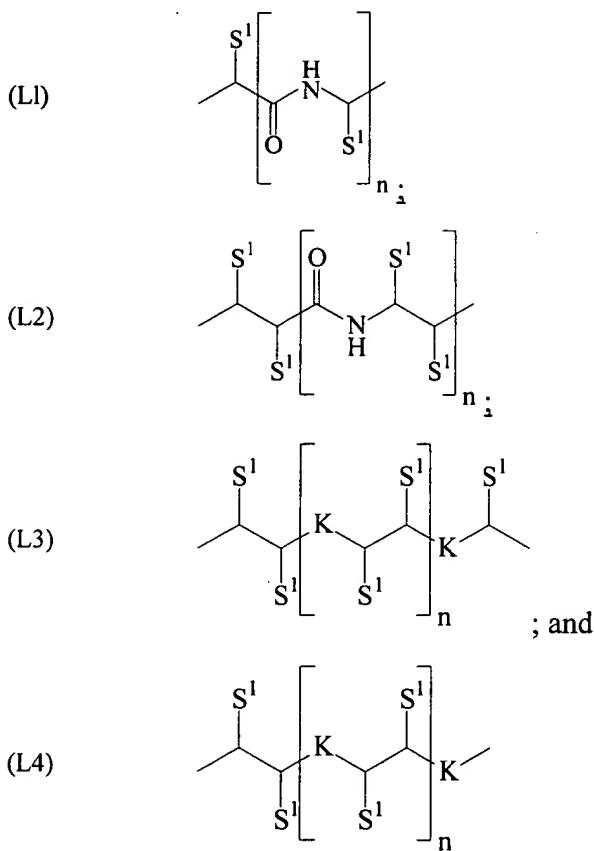
9. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core cyclic peptide of aspartocin.

10. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core antibiotic of aspartocin.

11. (Original) The antimicrobial sulfonamide derivative of Claim 4 in which m is 1.

12. (Original) The antimicrobial sulfonamide derivative of Claim 4 in which R<sup>1</sup> and R<sup>4</sup> are hydrogen.

13. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which L is selected from the group consisting of:



or a pharmaceutically acceptable salt or hydrate thereof, wherein:

n is 0, 1, 2 or 3;

each S<sup>1</sup> is independently selected from the group consisting of hydrogen, (C<sub>1</sub>-C<sub>10</sub>) alkyl optionally substituted with one or more of the same or different R<sup>5</sup> groups, (C<sub>1</sub>-C<sub>10</sub>) heteroalkyl optionally substituted with one or more of the same or different R<sup>5</sup> groups, (C<sub>5</sub>-C<sub>10</sub>) aryl optionally substituted with one or more of the same or different R<sup>5</sup> groups, (C<sub>5</sub>-C<sub>15</sub>) arylaryl optionally substituted with one or more of the same or different R<sup>5</sup> groups, (C<sub>5</sub>-C<sub>15</sub>) biaryl optionally substituted with one or more of the same or different R<sup>5</sup> groups, five to ten membered heteroaryl optionally substituted with one or more of the same or different R<sup>5</sup> groups, (C<sub>6</sub>-C<sub>16</sub>) arylalkyl optionally substituted with one or more of the same or different R<sup>5</sup> groups and six to sixteen membered heteroarylalkyl optionally substituted with one or more of the same or different R<sup>5</sup> groups;

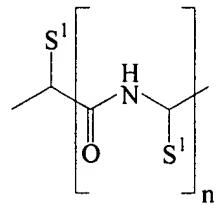
each R<sup>5</sup> is independently selected from the group consisting of -OR<sup>6</sup>, -SR<sup>6</sup>, -NR<sup>6</sup>R<sup>6</sup>, -CN, -NO<sub>2</sub>, -N<sub>3</sub>, -C(O)OR<sup>6</sup>, -C(O)NR<sup>6</sup>R<sup>6</sup>, -C(S)NR<sup>6</sup>R<sup>6</sup>, -C(NR<sup>6</sup>)NR<sup>6</sup>R<sup>6</sup>, -CHO, -R<sup>6</sup>CO, -SO<sub>2</sub>R<sup>6</sup>, -SOR<sup>6</sup>, -PO(OR<sup>6</sup>)<sub>2</sub>, -PO(OR<sup>6</sup>), -CO<sub>2</sub>H, -SO<sub>3</sub>H, -PO<sub>3</sub>H, halogen and trihalomethyl;

each R<sup>6</sup> is independently selected from the group consisting of hydrogen, (C<sub>1</sub>-C<sub>6</sub>) alkyl, (C<sub>5</sub>-C<sub>10</sub>) aryl, five to sixteen membered heteroaryl, (C<sub>6</sub>-C<sub>16</sub>) arylalkyl and six to sixteen membered heteroarylalkyl; and

each K is independently selected from the group consisting of oxygen, nitrogen and sulfur.

14. (Original) The antimicrobial sulfonamide of Claim 13 in which each S<sup>1</sup> is independently a side-chain of a genetically encoded  $\alpha$ -amino acid.

15. (Previously Presented) The antimicrobial sulfonamide of Claim 13 in which L is:



16. (Original) The antimicrobial sulfonamide derivative of Claim 15 in which each S<sup>1</sup> is independently a side-chain of a genetically encoded  $\alpha$ -amino acid.

17. (Original) The antimicrobial sulfonamide derivative of Claim 15 in which n is 0.

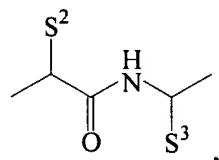
18. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 17 in which S<sup>1</sup> is hydrogen, Y is decan-yl and R is the core cyclic peptide of aspartocin.

19. (Original) The antimicrobial sulfonamide derivative of Claim 17 in which S<sup>1</sup> is -CH<sub>2</sub>-CO<sub>2</sub>H, -CH<sub>2</sub>-CH<sub>2</sub>-CO<sub>2</sub>H, -C(OH)H-CONH<sub>2</sub>, -CH<sub>2</sub>-CONH<sub>2</sub> or -CH<sub>2</sub>-CH<sub>2</sub>-CONH<sub>2</sub> or a salt or hydrate thereof.

20. (Original) The antimicrobial sulfonamide derivative of Claim 17 in which S<sup>1</sup> is -CH<sub>2</sub>-indol-2-yl or -CH<sub>2</sub>-phenyl.

21. (Cancelled)

22. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 13 in which L is:



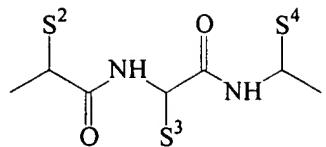
wherein S<sup>2</sup> and S<sup>3</sup> are each independently a side chain of a genetically encoded  $\alpha$ -amino acid.

23. (Cancelled)

24. (Original) The antimicrobial sulfonamide derivative of Claim 22 in which S<sup>2</sup> is hydrogen, -CH<sub>2</sub>-indol-2-yl, -CH<sub>2</sub>-CONH<sub>2</sub> or -CH<sub>2</sub>-CH<sub>2</sub>-CONH<sub>2</sub> and S<sup>3</sup> is -CH<sub>2</sub>-CO<sub>2</sub>H, -CH<sub>2</sub>-CH<sub>2</sub>-CO<sub>2</sub>H or a salt or hydrate thereof.

25. (Original) The antimicrobial sulfonamide derivative of Claim 22 in which S<sup>2</sup> is -CH<sub>2</sub>-CO<sub>2</sub>H, -CH<sub>2</sub>-CH<sub>2</sub>-CO<sub>2</sub>H or a salt or hydrate thereof and S<sup>3</sup> is -C(OH)H-CONH<sub>2</sub>.

26. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 13 in which L is:



wherein  $S^2$ ,  $S^3$ , and  $S^4$  are each independently a side chain of a genetically encoded  $\alpha$ -amino acid.

27. (Cancelled)

28. (Original) The antimicrobial sulfonamide derivative of Claim 26 in which  $S^2$  is  $-CH_2\text{-indol-2-yl}$ ,  $S^3$  is  $-CH_2\text{CONH}_2$  or  $-CH_2\text{-CH}_2\text{-CONH}_2$  and  $S^4$  is  $-CH_2\text{CO}_2\text{H}$ ,  $-CH_2\text{-CH}_2\text{-CO}_2\text{H}$  or a salt or hydrate thereof.

29. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 26 in which  $S^2$  is  $-CH_2\text{-indol-2-yl}$ ,  $S^3$  is  $-CH_2\text{CO}_2\text{H}$ ,  $CH_2\text{-CH}_2\text{-CO}_2\text{H}$  or a salt or hydrate thereof and  ~~$S^4$~~   $S^4$  is  $-CH_2\text{CONH}_2$ ,  $-CH_2\text{-CH}_2\text{-CONH}_2$  or  $-C(OH)H\text{-CONH}_2$ .

30. (Original) The antimicrobial sulfonamide derivative of Claim 4 in which  $m$  is 0.

31. (Original) The antimicrobial sulfonamide derivative of Claim 30 in which  $R^4$  is hydrogen.

32. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 30 in which  $R$  is the core antibiotic of aspartocin.

33. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 30 in which  $R$  is the core cyclic peptide of aspartocin.

34. (Currently Amended) A pharmaceutical composition comprising an antimicrobial sulfonamide derivative according to ~~Claim 4~~ any one of Claims 1 to 5 and a pharmaceutically acceptable adjuvant, excipient, carrier or diluent.

35. (Currently Amended) A method for treating or preventing a microbial infection, said method comprising the step of administering to a subject a therapeutically effective amount of ~~a compound according to Claim 4 or a therapeutically effective amount of a~~ pharmaceutical composition according to Claim 34.

36. (Currently Amended) A method of inhibiting microbial growth, said method comprising the step of administering to a microbe an antimicrobially effective amount of ~~an antimicrobial sulfonamide derivative according to Claim 4 or an antimicrobially effective amount of~~ a pharmaceutical composition according to Claim 34.

37. (Previously Presented) A method for making an antimicrobial sulfonamide derivative comprising sulfonylating a core antibiotic or core cyclic peptide with a lipophilic sulfonyl derivative, thereby providing an antimicrobial sulfonamide derivative.

38. (Currently Amended) The method of Claim 37 in which the lipophilic sulfonyl derivative is ~~a~~ an activated lipophilic sulfonyl ester or a lipophilic sulfonyl halide.

39. (Original) The method of Claim 38 in which the activated lipophilic sulfonyl ester is a lipophilic hydroxybenzotriazole ester.

40. (Previously Presented) The method of Claim 38 in which the lipophilic sulfonyl halide is a lipophilic sulfonyl chloride.

41. (Currently Amended) A method for making an antimicrobial sulfonamide derivative comprising:

sulfonylating a linker with a lipophilic sulfonyl compound, thereby providing a lipophilic sulfonamide linker; and

covalently attaching the lipophilic sulfonamide linker to a core antibiotic or core cyclic peptide wherein said core cyclic peptide or core antibiotic is ~~not of polymyxin of an acidic lipopeptide antibiotic~~, thereby yielding an antimicrobial sulfonamide derivative.

42. (Previously Presented) A method for making an antimicrobial sulfonamide derivative comprising:

covalently attaching a linker to a core antibiotic or core cyclic peptide, thereby providing an linker core antibiotic or linker core cyclic peptide; and

sulfonylating the linker core antibiotic or linker core cyclic peptide with a lipophilic sulfonyl derivative, thereby yielding an antimicrobial sulfonamide derivative.

43. (New) A method for treating or preventing a microbial infection, said method comprising the step of administering to a subject a therapeutically effective amount of an antimicrobial sulfonamide derivative according to any one of Claims 1 to 5.

44. (New) The method of Claim 43 in which the core cyclic peptide is aspartocin.

45. (New) The method of Claim 43 in which the core antibiotic is aspartocin.

46. (New) A method of inhibiting microbial growth, said method comprising the step of administering to a microbe an antimicrobially effective amount of an antimicrobial sulfonamide derivative according to any one of Claims 1 to 5.

47. (New) The method of Claim 46 in which the core cyclic peptide is aspartocin.

48. (New) The method of Claim 46 in which the core antibiotic is aspartocin.